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From: Rusyn, Ivan
Sent: Mon 3/9/2015 8:42:51 AM
Subject: section 4 data in relation to MAL, DZN and GLY human and animal evidence

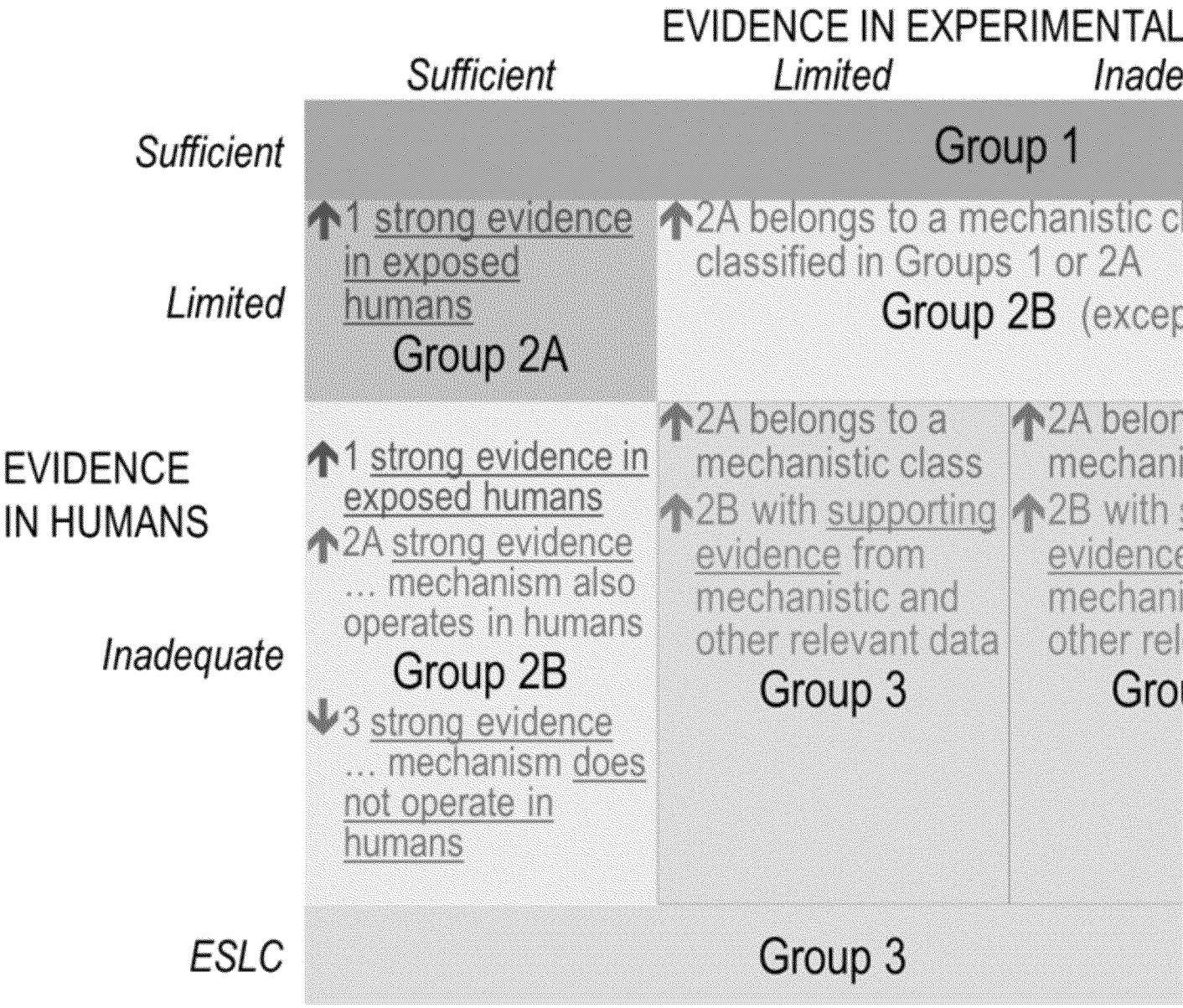
I would like to convene Group 4 downstairs in the first coffee break to discuss the information below.

Just to make sure we are all on the same page. Below are the evaluations from Groups 2 and 3 and the IARC matrix to get us to understand where our conclusions fit.

MAL: Human – Limited; Animal – sufficient → 2A; Group 4 evidence is strong to support carcinogenesis and we have data to show that the mechanisms can operate in humans, so we support the classification in 2A

DZN: Human – Limited; Animal – Inadequate (only one study) → 2B. Group 4 concludes that there is strong evidence for genotoxicity and oxidative stress and that these mechanisms can operate in humans. So we may consider upgrade to 2A.

GLY: Human – Limited; Animal – Limited → 2B. I have questions on the "limited" in animals as there are 2 studies showing significant effect… Nonetheless, Group 4 concludes that there is strong evidence for genotoxicity and oxidative stress and that these mechanisms can operate in humans. So we may consider upgrade to 2A.


EVIDENCE IN EXPERIMENTAL
Sufficient
Limited
Inade
EVIDENCE IN HUMANS
Sufficient
Limited
Inadequate
ESLC
Group 1
↑1 strong evidence in exposed humans
Group 2A
↑2A belongs to a mechanistic cl
classified in Groups 1 or 2A
Group 2B (excep
↑1 strong evidence in exposed humans
↑2A strong evidence … mechanism also operates in humans
Group 2B
↓3 strong evidence … mechanism does not operate in humans
↑2A belongs to a mechanistic class
↑2B with supporting evidence from mechanistic and other relevant data
Group 3
↑2A belon
mechani
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Group 3